

09428642

FILE 'HOME' ENTERED AT 14:04:16 ON 04 SEP 2001

=> file medline biosis embase capplus uspatfull

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	0.21	0.21

FILE 'MEDLINE' ENTERED AT 14:04:27 ON 04 SEP 2001

FILE 'BIOSIS' ENTERED AT 14:04:27 ON 04 SEP 2001  
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FILE 'USPATFULL' ENTERED AT 14:04:27 ON 04 SEP 2001  
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=> s androgen (p) receptor (p) slim3

=> dall ibib kwic

L1 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2001 ACS  
ACCESSION NUMBER: 2000:833264 CAPLUS  
DOCUMENT NUMBER: 134:13738  
TITLE: Use of SLIM3 for ligand screening  
INVENTOR(S): Schule, Roland; Muller, Judith  
PATENT ASSIGNEE(S): Schering A. G., Germany  
SOURCE: Jpn. Kokai Tokkyo Koho, 7 pp.  
CODEN: JKXXAF  
DOCUMENT TYPE: Patent  
LANGUAGE: Japanese  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 2000327587	A2	20001128	JP 1999-261593	19990916
EP 1058117	A1	20001206	EP 1999-250161	19990521
EP 1058117	B1	20010620		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
AT 202420	E	20010715	AT 1999-250161	19990521
PRIORITY APPLN. INFO.:			EP 1999-250161	A 19990521
AB Disclosed is the use of SLIM3 and its interaction with nucleus receptor protein, such as androgen receptor or				

.estrogen receptor .beta. subunit, for identification of ligands, antagonists and agonists.

ST SLIM3 protein androgen estrogen receptor ligand

IT Proteins, specific or class  
RL: ARG (Analytical reagent use); BSU (Biological study, unclassified); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)  
(SLIM3; protein SLIM3 for screening ligands or agonists and antagonists of nucleus receptor such as androgen receptor or estrogen receptor)

IT Drug screening  
Northern blot hybridization  
(protein SLIM3 for screening ligands or agonists and antagonists of nucleus receptor such as androgen receptor or estrogen receptor)

IT Ligands  
RL: ANT (Analyte); BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); PROC (Process); USES (Uses)  
(protein SLIM3 for screening ligands or agonists and antagonists of nucleus receptor such as androgen receptor or estrogen receptor)

IT Androgen receptors  
Estrogen receptors  
Nuclear receptors  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
(protein SLIM3 for screening ligands or agonists and antagonists of nucleus receptor such as androgen receptor or estrogen receptor)

IT cDNA  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(protein SLIM3 for screening ligands or agonists and antagonists of nucleus receptor such as androgen receptor or estrogen receptor)

=> s slim3

L2 12 SLIM3

=> dup rem 12

PROCESSING COMPLETED FOR L2

L3 6 DUP REM L2 (6 DUPLICATES REMOVED)

=> d l3 total ibib kwic

L3 ANSWER 1 OF 6 CAPLUS COPYRIGHT 2001 ACS  
ACCESSION NUMBER: 2000:772846 CAPLUS  
DOCUMENT NUMBER: 133:331185  
TITLE: Protein-protein interactions and their use in drug screening and disease diagnosis  
INVENTOR(S): Heichman, Karen; Bartel, Paul L.  
PATENT ASSIGNEE(S): Myriad Genetics, Inc., USA  
SOURCE: PCT Int. Appl., 87 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
-----	-----	-----	-----	-----

WO 2000065340 A1 20001102 WO 2000-US10651 20000421  
 W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU,  
 CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID,  
 IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV,  
 MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG,  
 SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM,  
 AZ, BY, KG, KZ, MD, RU, TJ, TM  
 RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,  
 DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,  
 CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

PRIORITY APPLN. INFO.: US 1999-130389 P 19990422  
 US 1999-140693 P 19990624  
 US 1999-156947 P 19990930  
 US 1999-163073 P 19991102  
 US 1999-168376 P 19991202  
 US 1999-168378 P 19991202

REFERENCE COUNT:

5

REFERENCE(S) :  
3rd

- (1) Ausubel; Short Protocols in Molecular Biology,  
ed, chapter 13 1995, P53
- (2) Gunster; Molecular Cell Biol 1997, V17(4), P2326 CAPLUS
- (3) Naya; Tissue-specific regulation of the insulin gene by a novel basic helix-loop-helix transcription factor 1995, V9, P1009 CAPLUS
- (4) Romanowski; Proc Natl Acad Sci 1996, V93, P10189 CAPLUS
- (5) Zilberman; Circ Res 1998, V82(5), P566 CAPLUS

IT Proteins, specific or class

RL: ANT (Analyte); ARG (Analytical reagent use); BPR (Biological process);  
 ANST (Analytical study); BIOL (Biological study); PROC (Process); USES (Uses)  
 (DRAL/FHL-2/**SLIM3**, complexes; protein-protein interactions and their use in drug screening and disease diagnosis)

L3 ANSWER 2 OF 6 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2000:833264 CAPLUS  
 DOCUMENT NUMBER: 134:13738  
 TITLE: Use of **SLIM3** for ligand screening  
 INVENTOR(S): Schule, Roland; Muller, Judith  
 PATENT ASSIGNEE(S): Schering A. G., Germany  
 SOURCE: Jpn. Kokai Tokkyo Koho, 7 pp.  
 CODEN: JKXXAF  
 DOCUMENT TYPE: Patent  
 LANGUAGE: Japanese  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 2000327587	A2	20001128	JP 1999-261593	19990916
EP 1058117	A1	20001206	EP 1999-250161	19990521
EP 1058117	B1	20010620		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
AT 202420	E	20010715	AT 1999-250161	19990521
EP 1999-250161 A 19990521				

PRIORITY APPLN. INFO.:

TI Use of **SLIM3** for ligand screening  
 AB Disclosed is the use of **SLIM3** and its interaction with nucleus receptor protein, such as androgen receptor or estrogen receptor .beta. subunit, for identification of ligands, antagonists and agonists.

ST **SLIM3** protein androgen estrogen receptor ligand

IT Proteins, specific or class

RL: ARG (Analytical reagent use); BSU (Biological study, unclassified); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study);

USES (Uses)  
(protein **SLIM3** for screening ligands or  
agonists and antagonists of nucleus receptor such as androgen receptor  
or estrogen receptor)

IT Drug screening  
Northern blot hybridization  
(protein **SLIM3** for screening ligands or agonists and  
antagonists of nucleus receptor such as androgen receptor or estrogen  
receptor)

IT Ligands  
RL: ANT (Analyte); BPR (Biological process); BSU (Biological study,  
unclassified); THU (Therapeutic use); ANST (Analytical study); BIOL  
(Biological study); PROC (Process); USES (Uses)  
(protein **SLIM3** for screening ligands or agonists and  
antagonists of nucleus receptor such as androgen receptor or estrogen  
receptor)

IT Androgen receptors  
Estrogen receptors  
Nuclear receptors  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL  
(Biological study); PROC (Process)  
(protein **SLIM3** for screening ligands or agonists and  
antagonists of nucleus receptor such as androgen receptor or estrogen  
receptor)

IT cDNA  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(protein **SLIM3** for screening ligands or agonists and  
antagonists of nucleus receptor such as androgen receptor or estrogen  
receptor)

L3 ANSWER 3 OF 6 MEDLINE DUPLICATE 1  
ACCESSION NUMBER: 2001015909 MEDLINE  
DOCUMENT NUMBER: 20459249 PubMed ID: 11003643  
TITLE: FHL2 (**SLIM3**) is not essential for cardiac  
development and function.  
AUTHOR: Chu P H; Bardwell W M; Gu Y; Ross J Jr; Chen J  
CORPORATE SOURCE: Department of Medicine, School of Medicine, University of  
California at San Diego, La Jolla, California 92093-0613,  
USA.  
SOURCE: MOLECULAR AND CELLULAR BIOLOGY, (2000 Oct) 20 (20) 7460-2.  
Journal code: NGY; 8109087. ISSN: 0270-7306.  
PUB. COUNTRY: United States  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 200010  
ENTRY DATE: Entered STN: 20010322  
Last Updated on STN: 20010322  
Entered Medline: 20001030  
TI FHL2 (**SLIM3**) is not essential for cardiac development and  
function.

L3 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2001 ACS  
ACCESSION NUMBER: 2000:70353 CAPLUS  
DOCUMENT NUMBER: 132:80381  
TITLE: Online data compression and error analysis using  
wavelet technology  
AUTHOR(S): Misra, Manish; Qin, S. Joe; Kumar, Shailesh; Seemann,  
Dick  
CORPORATE SOURCE: Dept. of Chemical Engineering, The University of  
Texas  
at Austin, Austin, TX, 78712, USA  
SOURCE: AIChE J. (2000), 46(1), 119-132  
CODEN: AICEAC; ISSN: 0001-1541  
PUBLISHER: American Institute of Chemical Engineers  
DOCUMENT TYPE: Journal

LANGUAGE: English  
REFERENCE COUNT: 16  
REFERENCE(S):  
(1) Bader, F; InTech 1987, V53  
(2) Bakshi, B; AIChE J 1996, V42, P477 CAPLUS  
(3) Benelli, D; The Radio and Electronic Engr 1980,  
V50, P29  
(13) Mah, R; Comp Chem Eng 1995, V19, P129 CAPLUS  
(15) Watson, M; Ind Eng Chem Res 1998, V37, P267  
CAPLUS  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

AB Wavelet representation of a signal is efficient for process data compression. An online compression algorithm based on Haar wavelets is proposed here. As a new data point arrives, the algorithm computes all the approxn. coeffs. and updates the multiresoln. tree before it preps.

to receive the next data point. An efficient bookkeeping and indexing scheme improves compression ratio more significantly than batch-mode wavelet compression. Reconstruction algorithms and historian format for this bookkeeping are developed. Various anal. results on the bounds on compression ratio and sum of the square error that can be achieved using this algorithm are derived. Exptl. evaluation over two sets of plant data shows that wavelet compression is superior to conventional interpolative methods (such as boxcar, backward slope, and **SLIM3**) in terms of quality of compression measured both in time and frequency domain and that the proposed online wavelet compression algorithm performs better than the batch-mode wavelet compression algorithm due to the efficient indexing and bookkeeping scheme. The online algorithm combines the high quality of compression of wavelet-based methods and online implementation of interpolative compression algorithms at the same time.

L3 ANSWER 5 OF 6 BIOSIS COPYRIGHT 2001 BIOSIS  
ACCESSION NUMBER: 1999:523458 BIOSIS  
DOCUMENT NUMBER: PREV199900523458  
TITLE: Developing cardiomyocytes selectively express the LIM protein DRAL/**SLIM3**.  
AUTHOR(S): Kong, Yanfeng (1); Bassel-Duby, Rhonda S.; Sanders-Williams, R.  
CORPORATE SOURCE: (1) Univ. Texas Southwestern Med. Cent., Dallas, TX USA  
SOURCE: Circulation, (Oct. 27, 1998) Vol. 98, No. 17 SUPPL., pp. I58.  
Meeting Info.: 71st Scientific Sessions of the American Heart Association Dallas, Texas, USA November 8-11, 1998  
The American Heart Association  
. ISSN: 0009-7322.  
DOCUMENT TYPE: Conference  
LANGUAGE: English  
TI Developing cardiomyocytes selectively express the LIM protein DRAL/**SLIM3**.  
IT . . .  
& Systems of Organisms  
cardiomyocyte: circulatory system, muscular system, selectivity  
IT Diseases  
dilated cardiomyopathy: heart disease  
IT Chemicals & Biochemicals  
DRAL/**SLIM3**; MNF-alpha: DNA-binding protein  
IT Alternate Indexing  
Cardiomyopathy, Congestive (MeSH)

L3 ANSWER 6 OF 6 MEDLINE  
ACCESSION NUMBER: 96354835 MEDLINE  
DOCUMENT NUMBER: 96354835 PubMed ID: 8753811

DUPLICATE 2

TITLE: Slim defines a novel family of LIM-proteins expressed in skeletal muscle.  
AUTHOR: Morgan M J; Madgwick A J  
CORPORATE SOURCE: Department of Orthodontics, Eastman Dental Institute, London, United Kingdom.  
SOURCE: BIOCHEMICAL AND BIOPHYSICAL RESEARCH COMMUNICATIONS, (1996 Aug 14) 225 (2) 632-8.  
Journal code: 9Y8; 0372516. ISSN: 0006-291X.  
PUB. COUNTRY: United States  
Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
OTHER SOURCE: GENBANK-U60115; GENBANK-U60116; GENBANK-U60117; GENBANK-U60118  
ENTRY MONTH: 199610  
ENTRY DATE: Entered STN: 19961022  
Last Updated on STN: 19980206  
Entered Medline: 19961010

AB . . . . . novel single zinc finger domain located in the N-terminal region.

Similar sequences to SLIM were identified and termed SLIM2 and SLIM3. The SLIM3 cDNA sequence was identified subsequently as a partial sequence of the of the LIM-protein DRAL. The number and spacing of. . .

=> log y

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
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\* \* \* \* \* \* \* \* \* \* \* \* \* \* \* \* STN Columbus \* \* \* \* \* \* \* \* \* \* \* \* \* \* \*

FILE 'HOME' ENTERED AT 14:09:49 ON 04 SEP 2001

=> file medline biosis embase caplus uspatfull

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	0.21	0.21

FILE 'MEDLINE' ENTERED AT 14:10:01 ON 04 SEP 2001

FILE 'BIOSIS' ENTERED AT 14:10:01 ON 04 SEP 2001  
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FILE 'USPATFULL' ENTERED AT 14:10:01 ON 04 SEP 2001  
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=> s androgen (p) receptor (p) dral

L1 8 ANDROGEN (P) RECEPTOR (P) DRAL

=> s androgen (p) receptor (p) fhl2

L2 9 ANDROGEN (P) RECEPTOR (P) FHL2

=> l1 or l2

L1 IS NOT A RECOGNIZED COMMAND

The previous command name entered was not recognized by the system.  
For a list of commands available to you in the current file, enter  
"HELP COMMANDS" at an arrow prompt (>).

=> s l1 or l2

L3 13 L1 OR L2

=> dup rem 13

PROCESSING COMPLETED FOR L3

L4 4 DUP REM L3 (9 DUPLICATES REMOVED)

=> d l4 total ibib kwic

L4 ANSWER 1 OF 4 MEDLINE	DUPLICATE 1
ACCESSION NUMBER: 2001103482 MEDLINE	
DOCUMENT NUMBER: 20458893 PubMed ID: 11001931	
TITLE: Alzheimer's disease-associated presenilin 2 interacts with	

AUTHOR: DRAL, an LIM-domain protein.  
CORPORATE SOURCE: Tanahashi H; Tabira T  
Division of Demyelinating Disease and Aging, National  
Institute of Neuroscience, 4-1-1 Ogawahigashi, Kodaira,  
Tokyo 187-8502, Japan.. tanahash@ncnp.go.jp  
SOURCE: HUMAN MOLECULAR GENETICS, (2000 Sep 22) 9 (15) 2281-9.  
Journal code: BRC. ISSN: 0964-6906.

PUB. COUNTRY: ENGLAND: United Kingdom  
Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200102

ENTRY DATE: Entered STN: 20010322  
Last Updated on STN: 20010322  
Entered Medline: 20010208

AB Using the yeast two-hybrid system, we screened for proteins interacting with presenilin 2 (PS2) and cloned **DRAL**. **DRAL** is an LIM-only protein containing four LIM domains and an N-terminal half LIM domain. Previously **DRAL** has been cloned as a co-activator of the **androgen receptor** and as a protein interacting with a DNA replication regulatory protein, hCDC47. Our yeast two-hybrid assay showed that **DRAL** interacted with a hydrophilic loop region (amino acids 269-298) in the endoproteolytic N-terminal fragment of PS2, but not that of. . . this region, R275A, T280A, Q282A, R284A, N285A, P287T, I288L, F289A and S296A, in PS2 abolished the binding. This suggests

that **DRAL** recognizes the PS2 structure specifically. The in vitro interaction was confirmed by affinity column assay and the physiological interactions between endogenous PS2 and **DRAL** by co-immunoprecipitation from human lung fibroblast MRC5 cells.

Furthermore, in PS2-overexpressing HEK293 cells, we found an increase in the amount of **DRAL** in the membrane fraction and an increase in the amount of **DRAL** that was co-immunoprecipitated with PS2. The potential role of **DRAL** in the cellular signaling suggests that **DRAL** functions as an adaptor protein that links PS2 to an intracellular signaling.

L4 ANSWER 2 OF 4 MEDLINE DUPLICATE 2  
ACCESSION NUMBER: 2000120800 MEDLINE  
DOCUMENT NUMBER: 20120800 PubMed ID: 10654935  
TITLE: **FHL2**, a novel tissue-specific coactivator of the **androgen receptor**.  
AUTHOR: Muller J M; Isele U; Metzger E; Rempel A; Moser M;  
Pscherer A; Breyer T; Holubarsch C; Buettner R; Schule R  
CORPORATE SOURCE: Universitats-Frauenklinik, Abteilung Frauenheilkunde und  
Geburtshilfe I, Klinikum der Universitat Freiburg,  
Breisacherstrasse 117, 79106 Freiburg, Germany.  
SOURCE: EMBO JOURNAL, (2000 Feb 1) 19 (3) 359-69.  
Journal code: EMB; 8208664. ISSN: 0261-4189.  
PUB. COUNTRY: ENGLAND: United Kingdom  
Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 200003  
ENTRY DATE: Entered STN: 20000327  
Last Updated on STN: 20000327  
Entered Medline: 20000310

TI **FHL2**, a novel tissue-specific coactivator of the **androgen receptor**.  
AB The control of target gene expression by nuclear receptors requires the recruitment of multiple cofactors. However, the exact mechanisms by which nuclear receptor-cofactor interactions result in tissue-specific gene regulation are unclear. Here we characterize a novel tissue-specific coactivator for the **androgen**

. receptor (AR), which is identical to a previously reported protein FHL2/DRAL with unknown function. In the adult, FHL2 is expressed in the myocardium of the heart and in the epithelial cells of the prostate, where it colocalizes with the AR in the nucleus. FHL2 contains a strong, autonomous transactivation function and binds specifically to the AR in vitro and in vivo. In an agonist- and AF-2-dependent manner FHL2 selectively increases the transcriptional activity of the AR, but not that of any other nuclear receptor. In addition, the transcription of the prostate-specific AR target gene probasin is coactivated by FHL2. Taken together, our data demonstrate that FHL2 is the first LIM-only coactivator of the AR with a unique tissue-specific expression pattern.

L4 ANSWER 3 OF 4 EMBASE COPYRIGHT 2001 ELSEVIER SCI. B.V.  
ACCESSION NUMBER: 2000437875 EMBASE  
TITLE: [Tissue specificity of molecular androgen action, crucial role of transcriptional cofactors].  
SPECIFICITE TISSULAIRE DE L'ACTION MOLECULAIRE DES ANDROGENES: ROLE DES COFACTEURS TRANSCRIPTIONNELS.  
AUTHOR: Gobinet J.; Jalaguier S.; Sultan C.  
CORPORATE SOURCE: J. Gobinet, Inst. Natl./la Sante/Recherche Med., INSERM U439, Pathol. Molec. des Recept. Nucleaires, 70 rue de Navacelles, F-34090 Montpellier, France  
SOURCE: References en Gynecologie Obstetrique, (2000) 7/4-5 (262-266).  
Refs: 47  
ISSN: 1244-8168 CODEN: RGEBE2  
COUNTRY: France  
DOCUMENT TYPE: Journal; General Review  
FILE SEGMENT: 003 Endocrinology  
LANGUAGE: French  
SUMMARY LANGUAGE: English; French  
AB Androgens participate in the development and maintenance of adult testis and prostate, and their action is mediated by the androgen receptor (AR). The specificity of AR action depends on the capacity of enzymatic cells to transform hormonal precursors into testosterone, particularly. . . These cofactors are able to modulate the transcriptional activity of AR either by augmentation or inhibition. Two recently isolated cofactors, FHL2 and PIAS1, seem to be good candidates for the control of AR action because of the specificity of their action and expression. FHL2, a 32 kDa protein, is an AR-specific coactivator whose expression pattern is restricted to prostate and myocardium. PIAS1, a 76 kDa protein, is an AR coactivator whose expression pattern is restricted to testis, particularly in Sertoli and Leydig cells. FHL2 is a potential regulator of gene expression in prostate and PIAS1 could be a testicular modulator of transcription.

L4 ANSWER 4 OF 4 MEDLINE DUPLICATE 3  
ACCESSION NUMBER: 2001022482 MEDLINE  
DOCUMENT NUMBER: 20481833 PubMed ID: 11027411  
TITLE: Expression of androgen receptor coregulatory proteins in prostate cancer and stromal-cell culture models.  
AUTHOR: Nessler-Menardi C; Jotova I; Culig Z; Eder I E; Putz T; Bartsch G; Klocker H  
CORPORATE SOURCE: Department of Urology, University of Innsbruck, Innsbruck, Austria.  
SOURCE: PROSTATE, (2000 Oct 1) 45 (2) 124-31.  
Journal code: PB4. ISSN: 0270-4137.  
PUB. COUNTRY: United States  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 200011

ENTRY DATE: Entered STN: 20010322  
Last Updated on STN: 20010322  
Entered Medline: 20001109

AB BACKGROUND: Androgen receptor (AR) transcriptional activity is modulated by cofactor proteins. They act as costimulators, corepressors, or bridging proteins, and a disbalanced expression leads to the altered activity of the AR in advanced prostate cancer. We investigated the expression of a series of steroid receptor cofactors in prostate cancer cell lines, including several LNCaP sublines, and in prostate stromal cells. METHODS: Expression of cofactors was analyzed by means of RT-PCR in PC-3, Du-145, LNCaP, three sublines of LNCaP established after long-term androgen deprivation, and two strains of primary prostate stroma cells. Expression in LNCaP and LNCaP-abl cells (which represented an advanced tumor) was expressed in all cells analyzed (AIB1, ARA54, ARA70, CBP, cyclin D1, Her2/neu/erbB2, BAG-1/M/L, SRC-1, SMRT, and TIF2). Only ARA55 and FHL2 mRNAs were not detected in all cells. ARA55 mRNA was absent in LNCaP cells, LNCaP sublines, and DU-145 cells; FHL2 was not expressed in LNCaP cells and its derivatives. The expression pattern was identical in LNCaP cells, and the long-term androgen ablated LNCaP sublines. Moreover, comparison of expression levels in LNCaP and LNCaP-abl cells revealed a slight reduction in LNCaP-abl cells but no gross differences.

CONCLUSIONS: Prostatic cells express a great number of steroid receptor cofactors. AR activity thus seems to be modulated in a very complex way in prostate cells.

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=> d his

(FILE 'HOME' ENTERED AT 14:09:49 ON 04 SEP 2001)

FILE 'MEDLINE, BIOSIS, EMBASE, CAPLUS, USPATFULL' ENTERED AT 14:10:01 ON 04 SEP 2001

L1 8 S ANDROGEN (P) RECEPTOR (P) DRAL  
L2 9 S ANDROGEN (P) RECEPTOR (P) FHL2  
L3 13 S L1 OR L2  
L4 4 DUP REM L3 (9 DUPLICATES REMOVED)

=> dup rem l1

PROCESSING COMPLETED FOR L1  
L5 2 DUP REM L1 (6 DUPLICATES REMOVED)

=> dup rem l2

PROCESSING COMPLETED FOR L2  
L6 3 DUP REM L2 (6 DUPLICATES REMOVED)

=> d 15 total ibib kwic

L5 ANSWER 1 OF 2 MEDLINE DUPLICATE 1  
ACCESSION NUMBER: 2001103482 MEDLINE  
DOCUMENT NUMBER: 20458893 PubMed ID: 11001931  
TITLE: Alzheimer's disease-associated presenilin 2 interacts with DRAL, an LIM-domain protein.  
AUTHOR: Tanahashi H; Tabira T  
CORPORATE SOURCE: Division of Demyelinating Disease and Aging, National Institute of Neuroscience, 4-1-1 Ogawahigashi, Kodaira, Tokyo 187-8502, Japan.. tanahash@ncnp.go.jp  
SOURCE: HUMAN MOLECULAR GENETICS, (2000 Sep 22) 9 (15) 2281-9.  
Journal code: BRC. ISSN: 0964-6906.  
PUB. COUNTRY: ENGLAND: United Kingdom

LANGUAGE: Journal; Article; (JOURNAL ARTICLE)  
FILE SEGMENT: English  
ENTRY MONTH: Priority Journals  
ENTRY DATE: 200102  
Entered STN: 20010322  
Last Updated on STN: 20010322  
Entered Medline: 20010208

AB Using the yeast two-hybrid system, we screened for proteins interacting with presenilin 2 (PS2) and cloned **DRAL**. **DRAL** is an LIM-only protein containing four LIM domains and an N-terminal half LIM domain. Previously **DRAL** has been cloned as a co-activator of the **androgen receptor** and as a protein interacting with a DNA replication regulatory protein, hCDC47. Our yeast two-hybrid assay showed that **DRAL** interacted with a hydrophilic loop region (amino acids 269-298) in the endoproteolytic N-terminal fragment of PS2, but not that of . . . this region, R275A, T280A, Q282A, R284A, N285A, P287T, I288L, F289A and S296A, in PS2 abolished the binding. This suggests

that **DRAL** recognizes the PS2 structure specifically. The in vitro interaction was confirmed by affinity column assay and the physiological interactions between endogenous PS2 and **DRAL** by co-immunoprecipitation from human lung fibroblast MRC5 cells.

Furthermore, in PS2-overexpressing HEK293 cells, we found an increase in the amount of **DRAL** in the membrane fraction and an increase in the amount of **DRAL** that was co-immunoprecipitated with PS2. The potential role of **DRAL** in the cellular signaling suggests that **DRAL** functions as an adaptor protein that links PS2 to an intracellular signaling.

L5 ANSWER 2 OF 2 MEDLINE DUPLICATE 2  
ACCESSION NUMBER: 2000120800 MEDLINE  
DOCUMENT NUMBER: 20120800 PubMed ID: 10654935  
TITLE: FHL2, a novel tissue-specific coactivator of the androgen receptor.  
AUTHOR: Muller J M; Isele U; Metzger E; Rempel A; Moser M;  
Pscherer

CORPORATE SOURCE: A; Breyer T; Holubarsch C; Buettner R; Schule R  
Universitats-Frauenklinik, Abteilung Frauenheilkunde und  
Geburtshilfe I, Klinikum der Universitat Freiburg,  
Breisacherstrasse 117, 79106 Freiburg, Germany.

SOURCE: EMBO JOURNAL, (2000 Feb 1) 19 (3) 359-69.  
Journal code: EMB; 8208664. ISSN: 0261-4189.

PUB. COUNTRY: ENGLAND: United Kingdom  
Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 200003  
ENTRY DATE: Entered STN: 20000327

Last Updated on STN: 20000327  
Entered Medline: 20000310

AB The control of target gene expression by nuclear **receptors** requires the recruitment of multiple cofactors. However, the exact mechanisms by which nuclear **receptor**-cofactor interactions result in tissue-specific gene regulation are unclear. Here we characterize a novel tissue-specific coactivator for the **androgen receptor** (AR), which is identical to a previously reported protein FHL2/**DRAL** with unknown function. In the adult, FHL2 is expressed in the myocardium of the heart and in the epithelial cells. . . agonist- and AF-2-dependent manner FHL2 selectively increases the transcriptional activity of the AR, but not that of any other nuclear **receptor**. In addition, the transcription of the prostate-specific AR target gene probasin is coactivated by FHL2. Taken together, our data demonstrate . . .

=> d.his

(FILE 'HOME' ENTERED AT 14:09:49 ON 04 SEP 2001)

FILE 'MEDLINE, BIOSIS, EMBASE, CAPLUS, USPATFULL' ENTERED AT 14:10:01 ON  
04 SEP 2001

L1 8 S ANDROGEN (P) RECEPTOR (P) DRAL  
L2 9 S ANDROGEN (P) RECEPTOR (P) FHL2  
L3 13 S L1 OR L2  
L4 4 DUP REM L3 (9 DUPLICATES REMOVED)  
L5 2 DUP REM L1 (6 DUPLICATES REMOVED)  
L6 3 DUP REM L2 (6 DUPLICATES REMOVED)

=> d 16 total ibib kwic

L6 ANSWER 1 OF 3 MEDLINE DUPLICATE 1  
ACCESSION NUMBER: 2000120800 MEDLINE  
DOCUMENT NUMBER: 20120800 PubMed ID: 10654935  
TITLE: **FHL2**, a novel tissue-specific coactivator of the  
androgen receptor.  
AUTHOR: Muller J M; Isele U; Metzger E; Rempel A; Moser M;  
Pscherer  
CORPORATE SOURCE: A; Breyer T; Holubarsch C; Buettner R; Schule R  
Universitats-Frauenklinik, Abteilung Frauenheilkunde und  
Geburtshilfe I, Klinikum der Universitat Freiburg,  
Breisacherstrasse 117, 79106 Freiburg, Germany.  
SOURCE: EMBO JOURNAL, (2000 Feb 1) 19 (3) 359-69.  
Journal code: EMB; 8208664. ISSN: 0261-4189.  
PUB. COUNTRY: ENGLAND: United Kingdom  
Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 200003  
ENTRY DATE: Entered STN: 20000327  
Last Updated on STN: 20000327  
Entered Medline: 20000310  
TI **FHL2**, a novel tissue-specific coactivator of the  
androgen receptor.  
AB The control of target gene expression by nuclear receptors  
requires the recruitment of multiple cofactors. However, the exact  
mechanisms by which nuclear receptor-cofactor interactions  
result in tissue-specific gene regulation are unclear. Here we  
characterize a novel tissue-specific coactivator for the androgen  
receptor (AR), which is identical to a previously reported protein  
**FHL2/DRAL** with unknown function. In the adult, **FHL2** is  
expressed in the myocardium of the heart and in the epithelial cells of  
the prostate, where it colocalizes with the AR in the nucleus.  
**FHL2** contains a strong, autonomous transactivation function and  
binds specifically to the AR in vitro and in vivo. In an agonist- and  
AF-2-dependent manner **FHL2** selectively increases the  
transcriptional activity of the AR, but not that of any other nuclear  
receptor. In addition, the transcription of the prostate-specific  
AR target gene probasin is coactivated by **FHL2**. Taken together,  
our data demonstrate that **FHL2** is the first LIM-only coactivator  
of the AR with a unique tissue-specific expression pattern.

L6 ANSWER 2 OF 3 EMBASE COPYRIGHT 2001 ELSEVIER SCI. B.V.  
ACCESSION NUMBER: 2000437875 EMBASE  
TITLE: [Tissue specificity of molecular androgen action, crucial  
role of transcriptional cofactors].  
SPECIFICITE TISSULAIRE DE L'ACTION MOLECULAIRE DES  
ANDROGENES: ROLE DES COFACTEURS TRANSCRIPTIONNELS.  
AUTHOR: Gobinet J.; Jalaguier S.; Sultan C.  
CORPORATE SOURCE: J. Gobinet, Inst. Natl./la Sante/Recherche Med., INSERM

SOURCE: U439, Pathol. Molec. des Recept. Nucleaires, 70 rue de Navacelles, F-34090 Montpellier, France  
References en Gynecologie Obstetrique, (2000) 7/4-5 (262-266).  
Refs: 47  
ISSN: 1244-8168 CODEN: RGEBE2

COUNTRY: France  
DOCUMENT TYPE: Journal; General Review  
FILE SEGMENT: 003 Endocrinology  
LANGUAGE: French  
SUMMARY LANGUAGE: English; French

AB **Androgens** participate in the development and maintenance of adult testis and prostate, and their action is mediated by the **androgen receptor** (AR). The specificity of AR action depends on the capacity of enzymatic cells to transform hormonal precursors into testosterone, particularly. . . These cofactors are able to modulate the transcriptional activity of AR either by augmentation or inhibition. Two recently isolated cofactors, **FHL2** and **PIAS1**, seem to be good candidates for the control of AR action because of the specificity of their action and expression. **FHL2**, a 32 kDa protein, is an AR-specific coactivator whose expression pattern is restricted to prostate and myocardium. **PIAS1**, a 76 kDa protein, is an AR coactivator whose expression pattern is restricted to testis, particularly in Sertoli and Leydig cells. **FHL2** is a potential regulator of gene expression in prostate and **PIAS1** could be a testicular modulator of transcription.

L6 ANSWER 3 OF 3 MEDLINE DUPLICATE 2  
ACCESSION NUMBER: 2001022482 MEDLINE  
DOCUMENT NUMBER: 20481833 PubMed ID: 11027411  
TITLE: Expression of androgen receptor coregulatory proteins in prostate cancer and stromal-cell culture models.  
AUTHOR: Nessler-Menardi C; Jotova I; Culig Z; Eder I E; Putz T; Bartsch G; Klocker H  
CORPORATE SOURCE: Department of Urology, University of Innsbruck, Innsbruck, Austria.  
SOURCE: PROSTATE, (2000 Oct 1) 45 (2) 124-31.  
PUB. COUNTRY: United States  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 200011  
ENTRY DATE: Entered STN: 20010322  
Last Updated on STN: 20010322  
Entered Medline: 20001109

AB BACKGROUND: **Androgen receptor** (AR) transcriptional activity is modulated by cofactor proteins. They act as costimulators, corepressors, or bridging proteins, and a disbalanced expression. . . to the altered activity of the AR in advanced prostate cancer. We investigated the expression of a series of steroid **receptor** cofactors in prostate cancer cell lines, including several LNCaP sublines, and in prostate stromal cells. METHODS: Expression of cofactors was analyzed by means of RT-PCR in PC-3, Du-145, LNCaP, three sublines of LNCaP established after long-term **androgen** deprivation, and two strains of primary prostate stroma cells. Expression in LNCaP and LNCaP-abl cells (which represented an advanced tumor. . . expressed in all cells analyzed (AIB1, ARA54, ARA70, CBP, cyclin D1, Her2/neu/erbB2, BAG-1/M/L, SRC-1, SMRT, and TIF2). Only ARA55 and **FHL2** mRNAs were not detected in all cells. ARA55 mRNA was absent in LNCaP cells, LNCaP sublines, and DU-145 cells; **FHL2** was not expressed in LNCaP cells and its derivatives. The expression pattern was identical in LNCaP cells, and the long-term **androgen** ablated LNCaP sublines.

Moreover, comparison of expression levels in LNCaP and LNCaP-abl cells revealed a slight reduction in LNCaP-abl cells but no gross differences.  
CONCLUSIONS: Prostatic cells express a great number of steroid receptor cofactors. AR activity thus seems to be modulated in a very complex way in prostate cells.

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COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	19.63	19.84

\* \* \* \* \*

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\* \* \* \* \*

STN INTERNATIONAL LOGOFF AT 14:13:57 ON 04 SEP 2001

DERWENT-ACC-NO: 2001-042441  
DERWENT-WEEK: 200143  
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TITLE: In vitro use of SLIM3 protein for binding to nuclear receptors, useful for identifying modulators of the androgen and estrogen-beta receptors

INVENTOR: MUELLER, J; SCHUELE, R

PATENT-ASSIGNEE: SCHERING AG [SCHD]

PRIORITY-DATA: 1999EP-0250161 (May 21, 1999)

PATENT-FAMILY:

PUB-NO	PUB-DATE	LANGUAGE	
PAGES	MAIN-IPC		
DE 59900132 G	July 26, 2001	N/A	000
	G01N 033/68		
EP 1058117 A1	December 6, 2000	G	009
	G01N 033/68		
JP 2000327587	November 28, 2000	N/A	007
	A61K 045/00		
A	June 20, 2001	G	000
	G01N 033/68		
EP 1058117 B1			

DESIGNATED-STATES: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK N  
L PT RO SE SI AL AT BE CH DE DK ES FI FR GB GR IE IT LI LT LU LV  
MC MK NL PT RO  
SE SI

APPLICATION-DATA:

PUB-NO	APPL-DESCRIPTOR	APPL-NO
APPL-DATE		
DE 59900132G	N/A	1999DE-0500132
May 21, 1999		
DE 59900132G	N/A	1999EP-0250161
May 21, 1999		
DE 59900132G	Based on	EP 1058117
N/A		
EP 1058117A1	N/A	1999EP-0250161
May 21, 1999		
JP2000327587A	N/A	1999JP-0261593
September 16, 1999		

EP 1058117B1  
May 21, 1999

N/A

1999EP-0250161

INT-CL\_(IPC): A61K038/00; A61K045/00 ; A61P005/26 ;  
A61P005/28 ;  
A61P005/30 ; A61P005/32 ; A61P043/00 ; C07K014/47 ;  
G01N033/53 ;  
G01N033/68

ABSTRACTED-PUB-NO: EP 1058117A

BASIC-ABSTRACT: NOVELTY - Extracorporeal use of the SLIM3 protein for binding to at least one of the nuclear proteins androgen receptor (AR) and estrogen beta receptor (ERb). All proteins may be in modified forms with deletion, substitution or insertion of up to 10 amino acids, provided that the function of the parent protein is retained.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

(1) use of amino acid sequences of SLIM3, encoded by a cDNA, for binding amino acids sequences of AR and ERb, also encoded by cDNAs, where optionally one or more of the cDNAs are modified but have at least 85% homology to sequences that encode the native proteins and encode a protein with the same function as the native protein; and

(2) extracorporeal method for identifying ligands that modulate interaction between SLIM3 and AR or ERb.

USE - Binding of SLIM3 to AR and ERb is used to identify ligands (agonists or antagonists) that modulate SLIM3-nuclear receptor interactions. These ligands are useful as therapeutic agents or as lead compounds for pharmaceutical development.

ADVANTAGE - Compared with known co-activators, SLIM3 is highly specific, i.e. it interacts with only AR and ERb.

ABSTRACTED-PUB-NO: EP 1058117B

EQUIVALENT-ABSTRACTS: NOVELTY - Extracorporeal use of the SLIM3 protein for binding to at least one of the nuclear proteins androgen receptor (AR) and estrogen beta receptor (ERb). All proteins may be in modified forms with deletion, substitution or insertion of up to 10 amino acids, provided that the function of the parent protein is retained.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

(1) use of amino acid sequences of SLIM3, encoded by a cDNA, for binding amino acids sequences of AR and ERb, also encoded by cDNAs, where optionally one or more of the cDNAs are modified but have at least 85% homology to sequences that encode the native proteins and encode a protein with the same function as the native protein; and

(2) extracorporeal method for identifying ligands that modulate interaction between SLIM3 and AR or ERb.

USE - Binding of SLIM3 to AR and ERb is used to identify ligands (agonists or antagonists) that modulate SLIM3-nuclear receptor interactions. These ligands are useful as therapeutic agents or as lead compounds for pharmaceutical development.

ADVANTAGE - Compared with known co-activators, SLIM3 is highly specific, i.e. it interacts with only AR and ERb.

CHOSEN-DRAWING: Dwg.0/0

TITLE-TERMS:

VITRO PROTEIN BIND NUCLEAR RECEPTOR USEFUL IDENTIFY MODULATE ANDROGENIC OESTROGEN BETA RECEPTOR

DERWENT-CLASS: B04 D16 S03

CPI-CODES: B04-C01G; B04-H01; B04-K01; B04-N04A; B11-C07B5; B11-C08E; B12-K04E;

D05-H09;

EPI-CODES: S03-E14H; S03-E14H4;

CHEMICAL-CODES:

Chemical Indexing M1 \*01\*

Fragmentation Code

M423 M430 M782 M905 N102 P831 Q233 Q505

Specfic Compounds

A00H3K A00H3D A00H3M

Chemical Indexing M1 \*02\*

Fragmentation Code

M423 M430 M782 M905 N102 P831 Q233 Q505

Specfic Compounds

A00H1K A00H1D A00H1M

Chemical Indexing M6 \*03\*

Fragmentation Code

M905 P611 P612 P621 P622 P831 Q233 Q505 R513 R515

R521 R614 R626 R627 R633

SECONDARY-ACC-NO:

CPI Secondary Accession Numbers: C2001-012379

Non-CPI Secondary Accession Numbers: N2001-031824